U.S. Patent Application No. 10/522,000 Amendment dated November 21, 2008

Reply to Office Action of August 21, 2008

AMENDMENTS TO THE CLAIMS:

This listing of claims will replace all prior versions, and listings, of claims in the application:

Listing of Claims:

- 1. (Previously presented) A labeled single chain antibody having a structure in which a heavy chain and a light chain of an antibody are directly crosslinked through a linker, wherein the linker is bound to a labeling substance, wherein the linker comprises an amino acid sequence that is recognizable by a biotin ligase, and wherein the biotin ligase binds the labeling substance to the linker.
- 2. (Previously presented) The labeled single chain antibody of claim 1, wherein the heavy chain and the light chain of the antibody are variable regions.
- 3. (Canceled).
- 4. (Canceled).
- 5. (Previously presented) A labeled single chain antibody having a structure in which a heavy chain and a light chain of an antibody are crosslinked through a linker, wherein the linker comprises a labeling substance, and wherein the labeling substance is incorporated as one part of the linker part of the antibody.
- 6. (Previously presented) The labeled single chain antibody of claim 5, having a structure in which the heavy chain and the light chain that are variable regions of the antibody are crosslinked through the linker.
- 7. (Previously presented) The labeled single chain antibody of claim 1, having a structure in which the heavy chain and the light chain of the antibody are crosslinked through the linker, wherein the labeling substance is biotin.

- 8. (Canceled)
- 9. (Previously presented) The labeled single chain antibody according to claim 1, which has a Kd value that is equivalent to a Kd value of a parental antibody and which is produced by a cell-free protein translation system using wheat embryo.

10-19. (Canceled)

- 20. (Previously presented) A labeled single chain antibody which has a Kd value that is equivalent to a Kd value of a parental antibody and is produced by a method for producing a labeled single chain antibody wherein DNA is subjected to transcription and translation, utilizing a wheat embryo-derived cell-free protein translation system in the presence of a labeling substance and in the presence of an enzyme that catalyzes a disulfide bond exchange reaction, and wherein the DNA comprises DNAs encoding a heavy chain and a light chain of an antibody having binding ability against a specific antigen, wherein the heavy chain and the light chain are linked through a DNA encoding a linker wherein the linker comprises an amino acid sequence that is recognizable by a biotin ligase, and wherein the biotin ligase binds the labeling substance to the linker.
- 21. (Currently amended) A method for producing an immobilized single chain antibody, wherein any one of the antibodies described hereunder is brought into contact with a reaction plate compartmentalized into a plurality of regions having on the surface thereof a substance that binds specifically with a labeling substance of the antibody:
- 1) a labeled single chain antibody having a structure in which a heavy chain and a light chain of the antibody are crosslinked through a linker, and the linker is bound to a labeling substance, wherein the linker comprises an amino acid sequence that is recognizable by a biotin ligase, and wherein the biotin ligase binds the labeling substance to the linker;

- 2) a labeled single chain antibody having a structure in which a heavy chain and a light chain that are variable regions of the antibody are crosslinked through a linker, and the linker is bound to a labeling substance, wherein the linker comprises an amino acid sequence that is recognizable by a biotin ligase, and wherein the biotin ligase binds the labeling substance to the linker;
- 3) a labeled single chain antibody having a structure in which a heavy chain and a light chain of the antibody are crosslinked through a linker, and the linker is bound to a labeling substance wherein the linker comprises an amino acid sequence that is recognizable by a biotin ligase, wherein the biotin ligase binds the labeling substance to the linker and, wherein the labeling substance is biotin; or
- 4) a labeled single chain antibody having a structure in which a heavy chain and a light chain that are variable regions of the antibody are crosslinked through a linker, and the linker is bound to a labeling substance wherein the linker comprises an amino acid sequence that is recognizable by a biotin ligase, wherein the biotin ligase binds the labeling substance to the linker and, wherein the labeling substance is biotin.
- 22. (Original) The method for producing an immobilized single chain antibody of claim 21, wherein two or more kinds of different immobilized single chain antibodies are immobilized on a reaction plate compartmentalized into a plurality of regions.
- 23. (Previously presented) The production method according to claim 21, wherein a labeling substance is biotin and a substance that binds specifically with the labeling substance is streptavidin.
- 24. (Previously presented) An immobilized single chain antibody prepared by the production method according to claim 21.

- 25. (Withdrawn) A method for analyzing an antigen-antibody reaction, wherein a test substance is brought into contact with the immobilized single chain antibody of claim 24, and binding ability of the test substance against the immobilized single chain antibody is analyzed.
- 26. (Withdrawn) A method for analyzing an antigen-antibody reaction, comprising the steps of:
- (1) preparing a labeled single chain antibody under conditions in which a disulfide bond of a single chain antibody is retained, comprising the step of the following (i) or (ii):
- (i) producing a labeled single chain antibody by subjecting a DNA, in which DNAs encoding a heavy chain and a light chain of an antibody having binding ability with a specific antigen are linked through a DNA encoding a linker comprising a nucleotide sequence that is capable of binding with a labeling substance in the presence of a specific enzyme after translation, to transcription and translation utilizing a wheat cell-free protein synthesis system in the presence of a specific enzyme; or
 - (ii) producing a labeled single chain antibody by subjecting a DNA, in which DNAs encoding a heavy chain and a light chain that are variable regions of an antibody having binding ability with a specific antigen are linked through a DNA encoding a linker comprising a nucleotide sequence that is capable of binding with a labeling substance in the presence of a specific enzyme after translation, to transcription and translation utilizing a wheat cell-free protein synthesis system in the presence of a specific enzyme;
- (2) preparing a substance (adapter substance) that binds specifically with a labeling substance of a labeled single chain antibody in a case where the labeling substance of the labeled single chain antibody is an immobilizing substance, comprising the steps of:

- (i) immobilizing a substance (adapter substance) that binds specifically with a labeling substance of a labeled single chain antibody to a reaction plate compartmentalized into a plurality of regions;
- (ii) removing a substance (adapter substance) that binds specifically with a labeling substance of a labeled single chain antibody that was not immobilized to the reaction plate in the preceding (i); and
- (iii) before and after the step of the preceding (i) or (ii), removing nonspecific adsorption from the reaction plate as appropriate;
- (3) preparing an immobilized labeled single chain antibody in a case where a labeling substance of the labeled single chain antibody is an immobilizing substance, comprising the steps of:
 - (i) adding a required amount of the labeling substance of the labeled single chain antibody prepared in (i) or (ii) of the above (1) onto a reaction plate compartmentalized into a plurality of regions having a substance (adapter substance) of (2) that binds specifically with the labeling substance of the labeled single chain antibody on the surface thereof, whereby to contact;
 - (ii) removing a labeled single chain antibody that was not immobilized to the substance (adapter substance) that binds specifically to the labeled single chain antibody on the reaction plate in the preceding (i); and
 - (iii) following the preceding step (ii), removing nonspecific adsorption from the reaction plate as appropriate;
- (4) preparing a labeled single chain antibody in a case where a labeling substance is a signal substance, comprising the steps of:

- (i) removing nonspecific adsorption from a reaction plate compartmentalized into a plurality of regions as appropriate; and
- (ii) adding a required amount of the labeling substance of the labeled single chain antibody prepared in (i) or (ii) of the above (1) onto the reaction plate;
- (5) adding a required amount of a test substance onto each reaction plate according to the above (3) or (4), and analyzing the binding ability of a labeled single chain antibody with the test substance; and
- (6) based on the binding ability result obtained in the above (5), qualitatively or quantitatively determining the interaction between the labeled single chain antibody and the test substance.
- 27. (Withdrawn) A reagent kit for measuring an antigen-antibody reaction, comprising a reagent to be used in the analysis method according to claim 25.
- 28. (Previously presented) An immobilized single chain antibody that has a Kd value that is equivalent to a Kd value of a parental antibody and that is produced by the method for producing an immobilized single chain antibody according to claim 21 utilizing a wheat embryo-derived cell-free protein translation system.
- 29. (Previously presented) A labeled single chain antibody which has a Kd value that is equivalent to a Kd value of a parental antibody and is produced by a method for producing a labeled single chain antibody wherein DNA is subjected to transcription and translation, utilizing a wheat embryo-derived cell-free protein translation system in the presence of a labeling substance and in the presence of an enzyme that catalyzes a disulfide bond exchange reaction, and wherein the DNA comprises DNAs encoding a heavy chain and a light chain of an antibody having binding ability against a specific antigen, wherein the heavy chain and the light chain are

linked through a DNA encoding a linker, wherein the linker comprises a labeling substance, and wherein the labeling substance is incorporated as one part of the linker part of the antibody.